## IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applica	ation of:	Senanayake et al.			
Serial N	No.:	Not yet assigned	Art Unit:	1621	
Filed:		Concurrently herewith	Examiner:	B. Davis	
For:		METHODS OF PREPARING DIDESMETHYLSIBUTRAMINE AND OTHER SIBUTRAMINE DERIVATIVES	Attorney D	ocket No.:	4821-522-999
		INFORMATION DISCLOSUE	RE STATEM	IENT	
P.O. Bo	ox 1450	or Patents 22313-1450			
Sir:					
subject	ed certain U.S. pate	dance with the duty of disclosure provision information which the Examiner may content application. It is requested that the Examination of the application.	isider materia	al to the exar	nination of the
1.	Enclosu	res accompanying this Information Disclo	sure Stateme	nt are:	
	la.	A list of all patents, publications, appl consideration by the office.	ications, or o	other informa	ation submitted for
	1b.	A legible copy of:			
		Each U.S. patent application publicati	on and U.S.	and foreign p	patent;
		Each publication or that portion which	n caused it to	be listed on	the PTO-1449;
		For each cited pending U.S. application the claims, and any drawing of the application caused it to be listed on the PTO-1449 incomes.	ition, or porti	on of the ap	plication which
		all other information or portion which	caused it to	be listed on	the PTO-1449.
	1c.	An English language copy of search reapplication or PCT International Search		a counterpa	art foreign
	1d.	Explanations of relevancy (ATTACH abstracts of the non-English language			nglish language
2.		This Information Disclosure Statement is  Within three months of the filing date continued prosecution application und	of a national		
		Within three months of the date of ent §1.491 in an international application;		ional stage a	s set forth in

		Before the mailing of the first Office action on the merits;
		Before the mailing of a first Office action after the filing of a request for continued examination under §1.114.
3.		This Information Disclosure Statement is filed under 37 C.F.R. §1.97(c) after the period specified in 37 C.F.R §1.97(b), but before the mailing date of any of a final action under 37 C.F.R. §1.113, a notice of allowance under 37 C.F.R. §1.311 or an action that otherwise closes prosecution in the application.
		(Check either Item 3a or 3b)
	3a.	☐ The Certification Statement in Item 5 below is applicable. Accordingly, no fee is required.
	3b.	☐ The \$180.00 fee set forth in 37 C.F.R. §1.17(p) in accordance with 37 C.F.R. §1.97(c) is: ☐ enclosed
		to be charged to Pennie & Edmonds LLP Deposit Account No. 16-1150.
		(Item 3b to be checked if any reference known for more than 3 months)
4.		This Information Disclosure Statement is filed under 37 C.F.R. §1.97(d) after the period specified in 37 C.F.R. §1.97(c), but on or before the date of payment of the issue fee.
		The \$180.00 fee set forth in 37 C.F.R. §1.17(p) is:  enclosed.  to be charged to Pennie & Edmonds LLP Deposit Account No. 16-1150.
	The Ce	rtification Statement in Item 5 below is applicable.
5.		Certification Statement (applicable if Item 3a or Item 4 is checked)
		(Check either Item 5a or 5b)
	5a.	☐ In accordance with 37 C.F.R. §1.97(e)(1), it is certified that each item of information contained in this Information Disclosure Statement was first cited in a communication from a foreign patent office in a counterpart foreign application not more than three months prior to the filing of this Information Disclosure Statement.
	5b.	In accordance with 37 C.F.R. §1.97(e)(2), it is certified that no item of information contained in this Information Disclosure Statement was cited in a communication from a foreign patent office in a counterpart foreign application and, to the knowledge of the undersigned after making reasonable inquiry, was known by any individual designated in 37 C.F.R. §1.56(c) more than three months prior to the filing of this Information Disclosure Statement.
6.	$\boxtimes$	This application is a continuation application under 37 C.F.R. §1.60 or §1.53(b) or (d).
		(Check appropriate Items 6a, 6b and/or 6c)
	6a.	A Petition to Withdraw from issue under 37 C.F.R. §1.313(b)(5) is concurrently filed herewith.

.

	6b.	Copies of publications listed on Form PTO-1449 from prior application Serial No. 10/120,503, filed on April 12, 2002, of which this application claims priority under 35 U.S.C. §120, are not being submitted pursuant to 37 C.F.R. §1.98(d).
	6c.	Copies of the publications listed on Form PTO-1449 were not previously cited in prior application Serial No. , filed on , and are provided herewith.
7.		This is a Supplemental Information Disclosure Statement. (Check Item 7a)
	7 <b>a</b> .	This Supplemental Information Disclosure Statement under 37 C.F.R. §1.97(f) supplements the Information Disclosure Statement filed on . A bona fide attempt was made to comply with 37 C.F.R. §1.98, but inadvertent omissions were made. These omissions have been corrected herein. Accordingly, additional time is requested so that this Supplemental Information Disclosure Statement can be considered as if properly filed on .
8.		In accordance with 37 C.F.R. §1.98, a concise explanation of what is presently understood to be the relevance of each non-English language publication is:
		(Check Item 8a, 8b, or 8c)
	8a.	satisfied because all non-English language publications were cited on the enclosed English language copy of the PCT International Search Report or the search report from a counterpart foreign application indicating the degree of relevance found by the foreign office.
	8b.	set forth in the application.
	8c.	enclosed as an attachment hereto.
9.	$\boxtimes$	The Commissioner is authorized to charge any additional fee required or credit any overpayment for this Information Disclosure Statement and/or Petition to Pennie & Edmonds LLP Deposit Account No. 16-1150.
10.		No admission is made that the information cited in this Statement is, or is considered to be, material to patentability nor a representation that a search has been made (other than a search report of a foreign counterpart application or PCT International Search Report if submitted herewith). 37 C.F.R. §§1.97(g) and (h).
		Respectfully submitted,
Date:	<u>Augus</u>	1 20, 2003 Dean L. Fanelli (Reg. No.)
		For: Anthony M. Insogna (Reg. No. 35,203)  PENNIE & EDMONDS LLP  1667 K St., N.W.  Washington, DC 20006  (202) 496-4400

,

.

## LIST OF REFERENCES CITED BY APPLICANT

(Use several sheets if necessary)

	ATTY DOCKET NO	APPLICATION NO		
	4821-414-999	10/120,503		
	APPLICANT			
C. Senanayake et al.				
	FILING DATE	GROUP		
	April 12, 2002	1621		

*EXAMINER		DOCUMENT NUMBER	DATE	S. PATENT DOCUMENTS	CLASS	SUBCLASS	FIUNG DATE
INITIAL	-		<del></del>		COSS	SUBCLASS	IF APPROPRIAT
	AA	6,127,363	10/2000	Doherty et al.			
	<del>                                     </del>	6,174,925	01/2001	Bailey et al.			
	AC	6,046,242	11/1998	Emmelmann			
	AD	5,795,880	08/1998	Svec et al.			
	AE	5,780,051	07/1998	Eswara et al.			
· · · · · · · · · · · · · · · · · · ·	AF	5,733,566	03/1998	Lewis			
	AG	5,719,283	02/1998	Bell et al.			
	AH	5,674,553	10/1997	Shinoda et al.			
	Al	5,639,476	06/1997	Oshlack et al.			
	AJ	5,591,767	01/1997	Mohr et al.			
	AK	5,552,429	09/1996	Wong et al.			
	AL	5,459,164	10/1995	Vargas			
	АМ	5,436,272	07/1995	Scheinbaum			
	AN	5,354,556	10/1994	Sparks et al.			
	AO	5,250,534	10/1993	Bell et al.			
	AP -	5,120,548	06/1992	McClelland et al.			
	AQ	5,104,899	04/1992	Young et al.			
	AR	5,073,543	12/1991	Marshall et al.			
	AS	5,068,440	11/1991	Jeffery et al.			
	АТ	5,059,595	10/1991	Le Grazie			
	AU	4,988,814	01/1991	Abou-Gharbia et al.			
	AV	4,939,175	07/1990	Ukai et al.			
	AW	4,929,629	05/1990	Jeffery			
	AX	4,871,774	10/1989	Rees			
	AY	4,816,488	03/1989	Rees			
	AZ	4,814,352	03/1989	Jeffery et al.			
	ВА	4,806,570	02/1989	Jeffery et al.			
	88	4,746,680	05/1988	Jeffery et al.			-
	ВС	4,552,828	11/1985	Toya et al.			
	BD	4,522,828	06/1985	Jeffery et al.			
	BE	4,008,719	02/1977	Theeuwes et al.			
	. BF	3,960,891	06/1976	Malen et al.		-	

BG	3,916,899	11/1975	Theeuwes et al.				
ВН	3,845,770	11/1974	Theeuwes et al.				
ВІ	3,598,123	08/1971	Zaffaroni				
ВЈ	3,536,809	10/1970	Applezweig				
вк	3,471,515	10/1969	Troxler et al.				
BL	3,155,670	11/1964	Janssen et al.				_
ВМ	3,155,669	11/1964	Janssen et al.				
		FORE	EIGN PATENT DOCUMENTS				
	DOCUMENT NUMBER	DATE	COUNTRY	CLASS	SUBCLASS	TRANSLA	
BN	WO 00/56322	09/2000	PCT			YES	NO.
ВО	WO 00/56321	09/2000	РСТ				
BP BP	WO 00/56318	09/2000	PCT				
BQ	WO 00/56315	09/2000	PCT				
BR	WO 00/56314	09/2000	PCT				
BS	WO 00/56309	09/2000	PCT				
ВТ	WO 00/56306	09/2000	PCT		•		
BU	WO 00/56151	09/2000	PCT				
BV	WO 00/56150	09/2000	PCT				
BW	WO 00/56149	09/2000	PCT				
BX	WO 00/56148	09/2000	PCT				
BY	WO 00/54765	09/2000	PCT				
вz	WO 00/32182	11/1998	PCT				
CA	WO 99/33450	07/1999	PCT				
СВ	WO 98/13034	04/1998	PCT				
CC	WO 98/13033	04/1998	PCT				
CD	WO 98/11884	03/1998	PCT				
CE	WO 98/06722	02/1998	PCT				
CF	EP 0 781 561 A1	07/1997	EPO				
CG	WO 97/20810	06/1997	PCT				
СН	WO 97/03675	02/1997	PCT				
CI	WO 95/21615	08/1995	PCT				
C1	WO 95/20949	08/1995	PCT	_			
СК	WO 94/28902	12/1994	PCT				
CL	WO 94/00114	01/1994	PCT				
СМ	WO 94/00047	01/1994	PCT				
CN	WO 90/06110	06/1990	PCT				
со	WO 88/06444	09/1988	PCT				
СР	DE 32 12 682 A1	10/1982	Germany				
CQ	GB 2098602 A	11/1982	United Kingdom				

CR	EP 0 035 597	9/1981	Europe		<u></u>			<u> </u>	
	OTHER RE	FERENCES (	Including Author, Title, Date, P	ertinent Pages, Etc.)					
CS Baldessarini et al., <i>Life Sciences</i> <b>39</b> : 1765-1777, (1986).									
СТ	Buckett et al., "BTS 54 2: 167-172 (1988).	Buckett et al., "BTS 54 524-An Approach to Rapidly Acting Antidepressant," New Concepts in Depression 2: 167-172 (1988).							
си	Bucket et al., "The Pharmacology of Sibutramine Hydrochloride (BTS 54 524), A New Antidepressant which induces Rapid Noradrenergic Down-Regulation", <i>Prog. Neuro-Psychopharmacol. &amp; Biol. Psychiat.</i> 12: 575-584 (1988).								
cv	Buckett et al., "Sibutra	mine Hydroc	chloride," Drugs of the Fut	ure <b>13(8)</b> : 736-738	(1988).				
cw	Butler, D., Facile Cyclo	oalkylation of	Arylacetonitriles in Dimeth	yl Sulfoxide, J. Org.	Chem.,	36:1308-	1309 (19	971)	
сх	Canonne, P., et al., <i>Et</i> 21:155-58 (1980)	fet du Benze	ene Dans la Reaction de G	rignard sur les Nitr	iles, Tet	trahedron	Lett.,		
CY	Carstensen, J., Drug	Stability: Prin	nciples & Practice, 2d. Ed.	, pp. 379-80, Marce	el Dekk	er, NY, N	Y, (1995	——— 5).	
cz	Castello, R.A., et al., (1962)	Discoloration	n of Tablets Containing A	mines and Lactose	, Pham	n. Sci. 51	(2):106-	-108	
DA	Cheetham, S.C., et al. Uptake: Effect of Adm 737-743	, [ <sup>8</sup> H]Paroxe inistration of	tine Binding in Rat Frontal Various Antidepressant T	Cortex Strongly Coreatments, Neurop	orrelate. harmacı	s with [ <sup>9</sup> H] ology (199	<i>]5-HT</i> 93), 32((	8),	
ОВ	Cliffe et al., (S)-N-tert- A Selective Antagonis	Butyl-3-(4-(2- t at Presynap	-methoxyphenyl)-piperazir otic and Postsynaptic 5-HT	n-1-yl)-2-phenylprop na Receptors, Med.	oanamic Chem.,	<i>le [(S)-WA</i> 36:1509-	4 <i>Y-1001</i> 1510 (1!	1 <i>35]:</i> 993)	
DC	Diagnostic and Statist (1997).	ical Manual d	of Mental Disorders, Fourt	h Ed., American Ps	ychiatri	c Associa	ition,		
DD	Diagnostic and Statist	ical Manual d	of Mental Disorders, 3rd E	d., American Psych	niatric A	ssociation	n (1981)	).	
DE	Dreshfield et al., Enhancement of Fluoxetine-Dependent Increase of Extracellular Serotonin (5-HT) Level by (-)-Pindolol, an Antagonist at 5-HT <sub>1A</sub> Receptors, Neurochem. Res., 21(5):557-562 (1996)						∍ls		
DF	Evans et al. "Prevalence of Alzheimer's Disease in a Community Population of Older Persons." LA								
DG						J.A.M.A	١.		
DH			he apparent anti-depressa s, <b>85</b> : 7, page 31, No. 407		i (+) tra:	nylpromin	e in an		
DI	Goodman & Gilman, 1	The Pharmac	cological Basis of Therape	utics, 362-373, 404	(9 <sup>th</sup> ed.	. McGraw	- -Hill, 19	96)	
DJ			e Opioidergic System in the Pharmacol., Vol. 124, No.			sm of Acti	on of		
DK	Handbook of Pharmad	ceutical Exci	pients, 2 <sup>nd</sup> ed., Wade and	Willer eds., pp. 257	 '-259 (1	994)			
DL			iffects on Central 5-HT Fur harmacol. (1998), 125(2),		ne Hydro	ochloride	and Oth	ner	
DM	Hillver et al., (S)-5-Flu Med. Chem., 33:1541		xy-2-(dipropylamino)tetrali	x: A Putative 5-HT,	 <sub>^</sub> Recep	otor Antag	gonist, J		
DN	J. Med. Chem., Vol. 3	6, No. 17, 25	540 (1993)						
DO	Introduction to Pharm	aceutical Do	sage Forms, 4th ed., Lea	& Febiger, Philadel	phia (19	<del></del> 985).	_		
DP	Jacques et al., "Enant	iomers, Race	emates and Resolutions,*	(Wiley-Interscience	, NY, 1	<del></del> 981).			
DQ			utical Sciences, <b>78</b> : 9: 695			<u> </u>			
DR	Jeffery et al., Synthes	is of Sibutrai	mine, A Novel Cyclobutyla J. Chem. Soc. Perkin. Tra	lkylamine Useful in		atment of	f Obesit	у,	

DS	King et al., "Clinical Pharmacology of Sibutramine Hydrochloride (BTS 54524) A New Antidepressant, in Healthy Volunteers," <i>Clinical Pharmac.</i> <b>26</b> : 607-611 (1989).
DΤ	Kula et al., "Effects of N-Substituted Phenyltetrahydropyridines on Cerebral High-Affinity Synatosomal Uptake of Dopamine and Other Monoamines in Several Mammalian Species," Life Sciences <b>34</b> (26): 2567-2575, (1984).
DU	Luscombe et al., The Contribution of Metabolites to the Rapid and Potent Down-Regulation of Rat Cortical β-Adrenoceptors by the Putative Antidepressant Sibutramine Hydrochloride, Neuropharmacology, Vol. 28, No. 2, (1989) pp 129-134
DV	The Merck Manual of Diagnosis and Therapy, ??th Ed., Merck & Co., Inc., Whitehouse Station, NJ, (1999).
DW	Middlemiss et al., Centrally Active 5-HT Receptor Agonists and Antagonists, Neurosci. and Biobehv. Rev., 16:75-82 (1992)
DX	Moreau et al., Behavioral Profile of the 5-HT <sub>1A</sub> Receptor Antagonist (S)-UH-301 in Rodents and Monkeys, Brain Res. Bull., 29:901-04 (1992)
DY	Nakada et al., An Enantioconvergent Route to (-)-Kainic Acid, Tetrahedron Lett., 38:857-860 (1997)
DZ	Physician's Desk Reference® 473-475 (53 <sup>rd</sup> ed., 1999)
EA	Physician's Desk Reference® 475-476 (53 <sup>rd</sup> ed., 1999)
EB	Physician's Desk Reference® 764-766 (53 <sup>rd</sup> ed., 1999)
EC	Physician's Desk Reference® 823-825 (53 <sup>rd</sup> ed., 1999)
ED	Physician's Desk Reference® 978-979 (53 <sup>rd</sup> ed., 1999)
EE	Physician's Desk Reference® 1054-1056 (53 <sup>rd</sup> ed., 1999)
EF	Physician's Desk Reference® 1332-1334 (53 <sup>rd</sup> ed., 1999)
EG	Physician's Desk Reference® 1369-1370 (53 <sup>rd</sup> ed., 1999)
EH	Physician's Desk Reference® 1432-1436 (53rd ed., 1999)
ΕI	Physician's Desk Reference® 1494-1498 (53 <sup>rd</sup> ed., 1999)
EJ	Physician's Desk Reference® 1641-1645 (53rd ed., 1999)
EK	Physician's Desk Reference® 2004-2009 (53 <sup>rd</sup> ed., 1999)
EL	Physician's Desk Reference® 2075-2078 (53rd ed., 1999)
EM	Physician's Desk Reference® 2190-2192 (53 <sup>rd</sup> ed., 1999)
EN	Physician's Desk Reference® 2367-2368 (53 <sup>rd</sup> ed., 1999)
EO	Physician's Desk Reference® 2396-2399 (53 <sup>rd</sup> ed., 1999)
EP	Physician's Desk Reference® 2490-2493 (53 <sup>rd</sup> ed., 1999)
EQ	Physician's Desk Reference® 2516-2521 (53rd ed., 1999)
ER	Physician's Desk Reference® 2688-2691 (53 <sup>rd</sup> ed., 1999)
ES	Physician's Desk Reference® 2701-2704 (53rd ed., 1999)
ET	Physician's Desk Reference® 2720-2726 (53 <sup>rd</sup> ed., 1999)
EU	Physician's Desk Reference® 2735-2736 (53 <sup>rd</sup> ed., 1999)
EV	Physician's Desk Reference® 2886-2888 (53 <sup>rd</sup> ed., 1999)
EW	Physician's Desk Reference® 2908-2910 (53 <sup>rd</sup> ed., 1999)
EX	Physician's Desk Reference® 3092-3094 (53 <sup>rd</sup> ed., 1999)
EY	Physician's Desk Reference® 3101-3104 (53 <sup>rd</sup> ed., 1999)

E	EZ	Physician's Desk Reference® 3224-3225 (53	<sup>rd</sup> ed., 1999)		
F	FA Physician's Desk Reference® 3267-3272 (53 <sup>rd</sup> ed., 1999)				
F	FB	Physician's Desk Reference® 3307-3309 (53 <sup>rd</sup> ed., 1999)			
F	FC	Physician's Desk Reference® 3383-3384 (53	<sup>rd</sup> ed., 1999)		
F	FD	Physician's Desk Reference® 2520 (52 <sup>™</sup> ed.,	1998)		
F	FE	Physician's Desk Reference® 2958 (52 <sup>™</sup> ed.,	1998)		
F	FF	Remingtons: The Practice of TheScience and Pharmacy, 19th ed., Gennaro, ed., p. 1625 (1995)			
F	FG	Remington's Pharmaceutical Sciences, 18th ed., Mack Publishing, Easton PA (1990).			
F	FH	Remington's Pharmaceutical Sciences, 16th ed., Mack Publishing, Easton PA (1980).			
	Stock, M.J., Sibutramine: A Review of the Pharmacology of a Novel Anti-Obesity Agent, Int'l J. Obesity, 21(Supp. 1):S25-S29 (1997)				
	FJ	U.S. Pharmocopia (USP) SP (XXI)/NF (XVI)			
F	FK	Wilen et al., Tetrahedron, 2725-36 33(21) (1977).			
1	FL	Wilen, S.H., Tables of Resolving Agents and Optical Resolutions 268 (E.L. Eliel ed., Univ. of Notre Dame Press, Notre Dame, IN, 1972).			
EXAMINER			DATE CONSIDERED		

\*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.